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Atty Dkt. No. STAN-276  
USSN: 10/713,674

<b>CERTIFICATE OF MAILING</b>			
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<b>REQUEST FOR CORRECTED FILING RECEIPT</b>  Address to:  Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450		Attorney Docket	STAN-276
		Applicant	LIN, KEN Y.
		Serial Number	10/713,674
		Filing Date	November 13, 2003
		Group Art Unit	1743
		Examiner Name	To Be Assigned
		Title: "METHODS FOR DETECTING ASYMMETRIC DIMETHYLARGININE IN A BIOLOGICAL SAMPLE"	

Sir:

A filing receipt for the above-identified patent application has been issued by the U.S. Patent and Trademark Office (copy attached) and has been found to contain the following error(s):

- (1) Please correct the filing date to read "November 13, 2003" as indicated on the application.

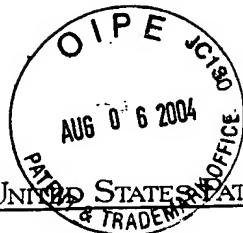
If for any reason a fee is found to be necessary, the Commissioner is authorized to charge such fee to Deposit Account No. 50-0815.

Respectfully submitted,  
BOZICEVIC, FIELD & FRANCIS LLP

Date: Aug. 2, 2004

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10/713,674	06/24/2004	1743	493	STAN-276	4	14	4

11/13/2004

CONFIRMATION NO. 9855

## UPDATED FILING RECEIPT



\*OC000000013272507\*

24353  
BOZICEVIC, FIELD & FRANCIS LLP  
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Date Mailed: 07/19/2004

Receipt is acknowledged of this regular Patent Application. It will be considered in its order and you will be notified as to the results of the examination. Be sure to provide the U.S. APPLICATION NUMBER, FILING DATE, NAME OF APPLICANT, and TITLE OF INVENTION when inquiring about this application. Fees transmitted by check or draft are subject to collection. Please verify the accuracy of the data presented on this receipt. If an error is noted on this Filing Receipt, please write to the Office of Initial Patent Examination's Filing Receipt Corrections, facsimile number 703-746-9195. Please provide a copy of this Filing Receipt with the changes noted thereon. If you received a "Notice to File Missing Parts" for this application, please submit any corrections to this Filing Receipt with your reply to the Notice. When the USPTO processes the reply to the Notice, the USPTO will generate another Filing Receipt incorporating the requested corrections (if appropriate).

## Applicant(s)

Ken Y. Lin, Boston, MA;  
John Cooke, Palo Alto, CA;

## Domestic Priority data as claimed by applicant

This appln claims benefit of 60/426,677-11/15/2002

## Foreign Applications

If Required, Foreign Filing License Granted: 02/23/2004

Projected Publication Date: 10/28/2004

Non-Publication Request: No

Early Publication Request: No

\*\* SMALL ENTITY \*\*

## Title

Methods for detecting asymmetric dimethylarginine in a biological sample

07/26/04  
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Bozicovic, Field, &amp; Francis



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**METHODS FOR DETECTING ASYMMETRIC DIMETHYLARGININE IN A BIOLOGICAL SAMPLE****CROSS-REFERENCE**

- [0001] This application claims the benefit of U.S. Provisional Patent Application No. 60/426,677 filed November 15, 2002, which application is incorporated herein by reference in its entirety.

**STATEMENT REGARDING FEDERALLY SPONSORED RESEARCH**

- [0002] The U.S. government may have certain rights in this invention, pursuant to grant no. R01 HL-63685 awarded by the National Institutes of Health, the National Heart, Lung and Blood Institute.

**FIELD OF THE INVENTION**

- [0003] The present invention is in the field of assay methods, and in particular assay methods for asymmetric dimethylarginine.

**BACKGROUND OF THE INVENTION**

- [0004] Elevated asymmetric dimethylarginine (ADMA) levels have been observed in various conditions, including hypertension, dyslipidemia, hyperglycemia, hyperhomocysteinemia, and renal failure, and are believed to be one cause of endothelial dysfunction in these conditions. Elevated plasma ADMA concentrations are also associated with an increased risk of cardiovascular disease. As an endogenous inhibitor of nitric oxide synthase, ADMA reduces nitric oxide (NO) production. NO plays a vital part in the vascular homeostasis. Aside from being the most potent vasodilator, NO inhibits platelet aggregation, smooth muscle proliferation, and adhesion molecule expression, which all play a part in atherogenesis. Throughout the last few years, basic scientific investigation has revealed the mechanism whereby ADMA becomes elevated in patients with hypertension, hyperhomocysteinemia, hyperglycemia, hypercholesterolemia, and tobacco exposure.
- [0005] Nevertheless, the field of ADMA is progressing slowly, mostly because of the laborious procedures required to quantify the molecule. A conclusive demonstration of